

PCT

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE



In re: Feng et al.

Serial Number: 10/525,410

Filed: January 26, 2005

For: **NOVEL BENZODIFURANIMIDAZOLINE  
AND BENZOFURANIMIDAZOLINE  
DERIVATIVES AND THEIR USE FOR  
THE TREATMENT OF GLAUCOMA**

CERTIFICATE OF MAILING

I hereby certify that this correspondence is being deposited with the United States Postal Service as First Class Mail in an envelope addressed to: MS Amendment, Commissioner for Patents, P. O. Box 1450, Alexandria, VA 22313-1450 on this date:

Date: August 2, 2005  
Jeannie Burke  
Jeannie Burke

**INFORMATION DISCLOSURE STATEMENT  
UNDER 37 CFR §1.97(b)(3)**

MS Amendment  
Commissioner for Patents  
P. O. Box 1450  
Alexandria VA 22313-1450

Sir:

Applicants submit the attached PTO Form 1449 and copies of sixty-three (63) references prior to the mailing date of a first Office Action on the merits in the above-referenced application.

Because the PTO has waived the requirement under 37 CFR 1.98(a)(2) to submit copies of each cited U.S. patent and patent application publication in all U.S. national patent applications filed after June 30, 2003, none are enclosed. Only copies of cited foreign patent documents and non-patent literature are enclosed.

Applicants request that the references listed in the accompanying PTO Form 1449 be considered during prosecution of the above-referenced application, and that these references appear among the "References Cited" on any patent issuing herefrom.

Respectfully submitted,  
ALCON

Date: 8/2/05

By:

Patrick M. Ryan  
Patrick M. Ryan  
Reg. No. 36,263

**ADDRESS FOR CORRESPONDENCE:**

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(817) 551-3066  
Atty. Docket No. 2345

Form PTO-1449 U.S. Dept. of Commerce

Patent & Trademark Office

LIST OF DOCUMENTS CITED BY APPLICANT

(Use several sheets if necessary)

ATTY. DOCKET NO.

2345 F US

SERIAL NO.

10/525,410

APPLICANT

Feng et al.

FILING DATE

January 26, 2005

GROUP

U.S. PATENT DOCUMENTS

INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
	AA	5,011,846	04/30/91	Gittos et al.	514	294	
	AB	5,290,781	03/01/94	Espino et al.	514	259	
	AC	5,494,928	02/27/96	Bös	514	415	
	AD	5,538,974	07/23/96	Ogawa et al.	514	253	
	AE	5,561,150	10/01/96	Wichmann	514	406	
	AF	5,571,833	11/05/96	Kruse et al.	514	414	
	AG	5,578,612	11/26/96	Macor et al.	514	323	
	AH	5,646,173	07/08/97	Bös et al.	514	411	
	AI	5,652,272	07/29/97	Ogawa et al.	514	652	
	AJ	5,693,654	12/02/97	Birch	514	323	
	AK	5,874,477	02/23/99	McConnell et al.	514	657	

FOREIGN PATENT DOCUMENTS

		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUBCLASS	TRANSLATION	
							YES	NO
	AL	CA 2,036,975	08/28/91	Canada				
	AM	EP 0 771 563 B1	01/22/03	Europe				
	AN	ES 323985	12/16/66	Spain				X
	AO	WO 92/00338	01/09/92	PCT				
	AP	WO 94/03162	02/17/94	PCT				

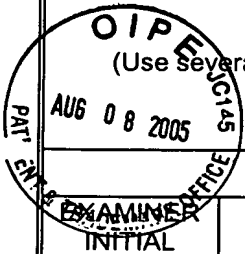
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)

AR	Boudier et al., "Structure Activity Relationships for Central and Peripheral Alpha Adrenergic Activities of Imidazoline Derivatives," <i>Life Sciences</i> , Vol. 17, pp. 377-386 (1975)
AS	Bowen et al., "Nonlinear regression using spreadsheets," <i>Trends in Pharmacological Sciences</i> , Vol. 16, pp. 413-417 (1995)
AT	Bradford et al., "A Rapid and Sensitive Method for the Quantitation of Microgram Quantities of Protein Utilizing the Principle of Protein-Dye Binding," <i>Analytical Biochemistry</i> , Vol. 72, pp. 248-254 (1976)

EXAMINER

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EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include a copy of this form with next communication to applicant.



Form PTO-1449 U.S. Dept. of Commerce  Patent & Trademark Office  <b>LIST OF DOCUMENTS CITED BY APPLICANT</b>  (Use several sheets if necessary)		<b>ATTY. DOCKET NO.</b> 2345F US		<b>SERIAL NO.</b> 10/525/410			
		<b>APPLICANT</b> Feng et al.					
		<b>FILING DATE</b> January 26, 2005		<b>GROUP</b> NYA			
U.S. PATENT DOCUMENTS							
EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUBCLASS	FILING DATE IF APPROPRIATE
	BA	5,891,131	04/06/99	Rajan et al.	606	5	
	BB	5,902,815	05/11/99	Olney et al.	514	285	
	BC	6,107,324	08/22/00	Behan et al.	514	406	
	BD	6,660,870	12/09/03	Ruskinko et al.	548	307.4	
	BE	6,664,286	12/16/03	May et al.	514	415	
	BF	6,696,476	02/24/04	Chen et al.	514	403	
	BG	6,806,285	10/19/04	May et al.	514	416	
	BH	6,884,816	04/26/05	May et al.	514	405	
	BI	2003/0181503	09/25/03	May et al.	514	406	
	BJ						
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FOREIGN PATENT DOCUMENTS							
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	BL	WO 97/35579	10/02/97	PCT			
	BM	WO 98/18458	05/07/98	PCT			
	BN	WO 98/31354	07/23/98	PCT			
	BO	WO 99/59499	11/25/99	PCT			
	BP	WO 00/12475	03/09/00	PCT			
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)							
	BR	Chambers et al., "Enantiospecific Synthesis and Pharmacological Evaluation of a Series of Super-Potent, Conformationally Restricted 5-HT <sub>2A/2C</sub> Receptor Agonists, <i>J. Med. Chem.</i> , Vol. 44, pp. 1003-1010 (2001)					
	BS	Chang et al., "Mechanism of the Ocular Hypotensive Action of Ketanserine," <i>J. of Ocular Pharmacology</i> , Vol. 1(2), pp. 137-147 (1985)					
	BT	De Vry et al., "Characterization of the Aminomethylchroman Derivative BAY X 3702 as a Highly Potent 5-Hydroxytryptamine <sub>1A</sub> Receptor Agonist," <i>J. of Pharmacology and Experimental Therapeutics</i> , Vol. 284 (3), pp. 1082-1094 (1998)					

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	CL	WO 00/12510	03/09/00	PCT		
	CM	WO 00/16761	03/30/00	PCT		
	CN	WO 00/35922	06/22/00	PCT		
	CO	WO 00/44753	08/03/00	PCT		
	CP	WO 01/40183	06/07/01	PCT		
<b>OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)</b>						
	CR	Fiorella et al., "The role of the 5-HT <sub>2A</sub> and 5-HT <sub>2C</sub> receptors in the stimulus effects of hallucinogenic drugs III: the mechanistic basis for supersensitivity to the LSD stimulus following serotonin depletion," <i>Psychopharmacology</i> , Vol. 121, pp. 364-372 (1995)				
	CS	Griffin et al., "Pharmacological Characterization of an FP Prostaglandin Receptor on Rat Vascular Smooth Muscle Cells (A7r5) Coupled to Phosphoinositide Turnover and Intracellular Calcium Mobilization," <i>J. of Pharmacology and Experimental Therapeutics</i> , Vol. 286(1), pp. 411-418 (1998)				
	CT	Gupta et al., "Therapeutic Potentials of 5-HT Receptor Modulators," <i>Indian J. of Pharmacology</i> , Vol. 26, pp. 94-107 (1994)				

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	DL	WO 01/70207	09/27/01	PCT				
	DM	WO 01/70223	09/27/01	PCT				
	DN	WO 01/70230	09/27/01	PCT				
	DO	WO 01/70701	09/27/01	PCT				
	DP	WO 01/70702	09/27/01	PCT				
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	DR	Hoyer et al., "VII. International Union of Pharmacology Classification of Receptors for 5-Hydroxytryptamine (Serotonin), <i>Pharmacological Reviews</i> , Vol. 46(2), pp. 157-203 (1994)						
	DS	Johnson et al., "Binding to the Serotonin 5-HT <sub>2</sub> Receptor by the Enantiomers of <sup>125</sup> I-DOI, <i>Neuropharmacology</i> , Vol. 26(12), pp. 1803-1806 (1987)						
	DT	Krootila et al., "Effect of Serotonin and its Antagonist (Ketanserin) in Intraocular Pressure in the Rabbit," <i>J. of Ocular Pharmacology</i> , Vol. 3(4), pp. 279-290 (1987)						

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	EN	WO 02/098400	12/12/02	PCT			
	EO	WO 02/098860	12/12/02	PCT			
	EP	WO 03/051291	06/26/03				
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)							
	ER	Lopez-Rodriguez, "Synthesis and Structure-Activity Relationships of a New Model of Arylpiperazines. 4. <sup>1</sup> -(ω-(4-Arylpiperazine-1-yl)alkyl)-3-(diphenylmethylene)-2,5-pyrrolidinediones and -3-(9H-fluoren-9-ylidene)-2,5-pyrrolidinediones: Study of the Steric Requirements of the Terminal Amide Fragment on 5-HT <sub>1A</sub> Affinity/Selectivity," J. Med. Chem., 42(1):36-49 (1999)					
	ES	Mallorga et al., "Characterization of serotonin receptors in the iris + ciliary body of the albino rabbit," <i>Current Eye Research</i> , Vol. 6(3), pp. 527-532 (1987)					
	ET	Martin et al., "The Structure and Signalling Properties of 5-HT Receptors: an endless diversity?" <i>Trends in Pharmaceutical Sciences</i> , Vol. 19, pp. 2-4 (1998)					

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	FN	WO 04/019874	03/11/04	PCT				
	FO	WO 04/028451	04/08/04	PCT				
	FP	WO 04/054572	07/01/04	PCT				
OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)								
	FR	Mano et al., "The Effect of Anplag (Sarpogelate HCL), New Selective 5-HT <sub>2</sub> Antagonist on Intraocular Pressure in Rabbits," <i>IOVS</i> , Vol. 36(4), S719 (1995)						
	FS	May et al., "A Novel and Selective 5-HT <sub>2</sub> Receptor Agonist with Ocular Hypotensive Activity: (S)-(+)-1-(2-Aminopropyl)-8,9-dihydropyrano[3,2-e]indole," <i>J. Med. Chem.</i> , Vol. 46, pp. 4188-4195 (2003)						
	FT	May et al., "Evaluation of the Ocular Hypotensive Response of Serotonin 5-HT <sub>1A</sub> and 5-HT <sub>2</sub> Receptor Ligands in Conscious Ocular Hypertensive Cynomolgus Monkeys," <i>J. of Pharmacology and Experimental Therapeutics</i> , Vol. 306(1), pp. 301-309 (2003)						

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	GL	WO 04/058725	07/15/04	PCT				
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<b>OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)</b>								
	GR	Monte et al, "Dihydrobenzofuran Analogues of Hallucinogens. 3 <sup>1</sup> Models of 4-Substituted (2,5-Dimethoxyphenyl)alkylamine Derivatives with Rigiddified Methoxy Groups," <i>J. Med. Chem.</i> , Vol. 39, pp. 2953-2961 (1996)						
	GS	Morin et al, "A Method to Measure Simultaneously Cyclic AMP and Inositol Phosphate Accumulation in Rat Brain Slices," <i>J. Of Neurochemistry</i> , Vol. 56(4), pp. 1114-1120 (1991)						
	GT	Osborne et al., "Do Beta-Adrenoceptors and Serotonin 5-HT <sub>1A</sub> Receptors Have Similar Functions in the Control of Intraocular Pressure in the Rabbit?", <i>Ophthalmologica</i> , Vol. 210, pp. 308-314 (1996)						

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							YES	NO
	HL							
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	HR	Osborne et al., "5-Hydroxytryptamine <sub>1A</sub> agonists: potential use in glaucoma. Evidence from animal studies," <i>Eye</i> , Vol. 14(38), pp. 454-463 (2000)						
	HS	Salomon et al., "Adenylate Cyclase Assay," <i>Advances in Cyclic Nucleotide Research</i> , Vol. 10, pp. 35-55 (1979)						
	HT	Salomon et al., "[2] Cellular Responsiveness to Hormones and Neurotransmitters: Conversion of [ <sup>3</sup> H]Adenine to [ <sup>3</sup> H]cAMP in Cell Monolayers, Cell Suspensions, and Tissue Slices," <i>Methods in Enzymology</i> , Vol. 195, pp. 22-28 (1991)						

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	IR	Schoeffter et al., "Inhibition of cAMP Accumulation Via Recombinant Human Serotonin 5-HT <sub>1A</sub> Receptors: Considerations on Receptor Effector Coupling across Systems," <i>Neuropharmacology</i> , Vol. 36(4/5), pp. 429-437 (1997)					
	IS	Sharif et al., "[ <sup>3</sup> H]AL-5848 ([ <sup>3</sup> H]9β-(+)-Fluprostenol). Carboxylic Acid of Travoprost (AL-6221), a Novel FP Prostaglandin o Study the Pharmacology and Autoradiographic Localization of the FP Receptor," <i>J. Pharm. Pharmacol.</i> Vol. 51: pp. 685-694 (1999)					
	IT	Takeda et al., "The Effect of Inplag. Novel Selective 5-HT <sub>2</sub> Antagonist on Intraocular Pressure in Glaucoma Patients," <i>IOVS</i> , Vol. 36(4), S734 (1995)					

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	JR	Tobin et al., "Evidence for the Presence of Serotonergic Nerves and Receptors in the Iris-Ciliary Body Complex of the Rabbit," <i>The J. of Neurosciences</i> , Vol. 8(10), pp. 3713-3721 (1988)						
	JS	Wang et al., "Effect of 5-methylurapidil, an $\alpha_{1a}$ -adrenergic antagonist and 5-hydroxytryptamine <sub>1a</sub> agonist on aqueous humor dynamics in monkeys and rabbits," <i>Current Eye Research</i> , Vol. 16(8), pp. 769-775 (1997)						
	JT	Wang et al., "Effect of Oxymetazoline on Aqueous Humor Dynamics and Ocular Blood Flow in Monkeys and Rabbits," <i>Archives of Ophthalmology</i> , Vol. 111, pp. 535-538 (1993)						

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OTHER PRIOR ART (Including Author, Title, Date, Pertinent Pages, Etc.)		
	KR	Wang et al., "Effect of p-MPPI Hydrochloride (p-MPPI) Applied Before 5-Methylurapidil (5-MU) on Intraocular Pressure (IOP) in Normal Monkeys," IVOS, 39(4), S488 (1998)
	KS	Zifa et al., "5-Hydroxytryptamine Receptors," <i>Pharmacological Reviews</i> , Vol. 44(3), pp. 401-458 (1992)
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EXAMINER

DATE CONSIDERED

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include a copy of this form with next communication to applicant.